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Entrez PubMed		1: Minier C, Lelong C, Djemel N, Rodet F, Tutundjian R, Favrel P, Mathieu M, Leboulenger F.  Related Articles, Links
		Expression and activity of a multixenobiotic resistance system in the Pacific
		oyster Crassostrea gigas.
		Mar Environ Res. 2002 Sep-Dec;54(3-5):455-9. PMID: 12408601 [PubMed - indexed for MEDLINE]
PubMed Services		2: Schwartz IF, Hershkovitz R, Iaina A, Gnessin E, Wollman Y,  Related Articles, Links
		Chernichowski T, Blum M, Levo Y, Schwartz D.  Garlic attenuates nitric oxide production in rat cardiac myocytes through
		inhibition of inducible nitric oxide synthase and the arginine transporter CAT-2
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		Vogensen SB, Jensen HS, Stensbol TB, Frydenvang K, Bang-Andersen B, Johansen TN, Egebjerg J, Krogsgaard-Larsen P.  Related Articles, Links
		Resolution, configurational assignment, and enantiopharmacology of
		2-amino-3-[3-hydroxy-5-(2-methyl-2H- tetrazol-5-yl)isoxazol-4-yl]propionic
Related		acid, a potent GluR3- and GluR4-preferring AMPA receptor agonist. Chirality. 2000 Nov;12(10):705-13.
Resourc	es	PMID: 11054828 [PubMed - indexed for MEDLINE]
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		Mutations in the organic cation/carnitine transporter OCTN2 in primary carnitine deficiency.
		Proc Natl Acad Sci U S A. 1999 Mar 2;96(5):2356-60.
		PMID: 10051646 [PubMed - indexed for MEDLINE]
		<b>5:</b> <u>Katiyar SK, Korman NJ, Mukhtar H, Agarwal R.</u> Related Articles, Links
		Protective effects of silymarin against photocarcinogenesis in a mouse skin model.
		J Natl Cancer Inst. 1997 Apr 16;89(8):556-66.
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		analogue, 1alpha-hydroxyvitamin D5.
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     Screening regulators of visceral fat accumulation by microarray gene
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     Matsuki, Yasushi; Iguchi, Haruhisa
     Sumitomo Chemical Co., Ltd., Japan
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     Arthur; Elashoff, Michael
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     Molecular toxicology modeling based on global changes in gene expression
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     Mendrick, Donna; Porter, Mark W.; Johnson, Kory R.; Castle, Arthur L.;
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     Method for detecting change in gene expression induced
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     Muramatsu, Masaaki; Wakao, Hiroshi; Wakao, Rika; Yano, Kazuhiro; Noguchi,
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Teruhisa; Suyama, Akira

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Helix Research Institute, Japan
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    PCT Int. Appl., 48 pp.
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      22090062 PubMed ID: 12096331
      [Improving the embryonic stem cell test (EST) by establishing molecular
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      endpoints of tissue specific development using murine embryonic stem
 cells
      (D3 cells)].
      Etablierung molekularer Endpunkte zur Weiterentwicklung des Embryonalen
      Stammzelltests (EST) mit embryonalen Stammzellen der Maus (Zelllinie
 D3).
      Seiler Andrea; Visan Anke; Pohl Ingeborg; Genschow Elke; Buesen Roland;
 ΑU
      Spielmann Horst
 CS
      Zentralstelle zur Erfassung und Bewertung von Ersatz- und
      Erganzungsmethoden zum Tierversuch (ZEBET), Bundesinstitut fur
      gesundheitlichen Verbraucherschutz und Veterinarmedizin (BgVV), D-Berlin,
      Germany.. seiler.zebet@bgvv.de
 SO
      ALTEX, (2002) 19 Suppl 1 55-63.
      Journal code: 100953980. ISSN: 0946-7785.
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      Priority Journals
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      PREV200100501809
      Molecular markers in embryonic stem cells.
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      zur Nieden, N. I. (1); Ruf, L. J.; Kempka, G.; Hildebrand, H.; Ahr, H. J.
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      (1) Research Toxicology, Bayer AG, 42096, Wuppertal:
 CS
      nicole.melzer.nm.@bayer.ag.de Germany
 SO
      Toxicology In Vitro, (August October, 2001) Vol. 15, No. 4-5, pp.
 455-461.
      print.
      ISSN: 0887-2333.
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      1999:487431 CAPLUS
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      131:112381
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      Gene expression fingerprint and use in compound
      screening for developing therapeutics
      Johnson, Paul H.; Ponte, Phyllis A.; Zajchowski, Deborah A.
 IN
      Schering Aktiengesellschaft, Germany
 PA
      PCT Int. Appl., 72 pp.
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      CODEN: PIXXD2
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      English
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     FILE 'BIOSIS, MEDLINE, EMBASE, LIFESCI, CAPLUS' ENTERED AT 11:40:39 ON
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1.2
L3
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              3 DUP REM L4 (2 DUPLICATES REMOVED)
L5
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MEDLINE IS NOT A RECOGNIZED COMMAND
The previous command name entered was not recognized by the system.
For a list of commands available to you in the current file, enter
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=> FIL MEDLINE
COST IN U.S. DOLLARS
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FULL ESTIMATED COST
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 FILE LAST UPDATED: 16 MAR 2003 (20030316/UP). FILE COVERS 1958 TO DATE.
On June 9, 2002, MEDLINE was reloaded. See HELP RLOAD for details.
MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the
MeSH 2003 vocabulary. See http://www.nlm.nih.gov/mesh/summ2003.html
 for a description on changes.
 This file contains CAS Registry Numbers for easy and accurate
 substance identification.
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L6 17 ("POWELL T J"/AU) AND 1990<=PY<=2000

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YOU HAVE REQUESTED DATA FROM 17 ANSWERS - CONTINUE? Y/(N):Y

- L6 ANSWER 1 OF 17 MEDLINE
- TI Chronic neurobehavioural effects of mercury poisoning on a group of Zulu chemical workers.
- L6 ANSWER 2 OF 17 MEDLINE
- TI Psychiatrists' referrals to self-help groups for people with mood disorders.
- L6 ANSWER 3 OF 17 MEDLINE
- TI A discrete subpopulation of dendritic cells transports apoptotic intestinal epithelial cells to T cell areas of mesenteric lymph nodes.
- L6 ANSWER 4 OF 17 MEDLINE
- TI Derivation of temporary emergency exposure limits (TEELs).
- L6 ANSWER 5 OF 17 MEDLINE
- TI Growth inhibition of psoriatic keratinocytes by quinazoline tyrosine kinase inhibitors.
- L6 ANSWER 6 OF 17 MEDLINE
- TI Recommended default methodology for analysis of airborne exposures to mixtures of chemicals in emergencies.
- L6 ANSWER 7 OF 17 MEDLINE
- ${\tt TI}$  Attitudes of AA contact persons toward group participation by persons with
  - a mental illness.
- L6 ANSWER 8 OF 17 MEDLINE
- TI Inhibition of platelet-derived growth factor-mediated signal transduction and tumor growth by N-[4-(trifluoromethyl)-phenyl]5-methylisoxazole-4-carboxamide.
- L6 ANSWER 9 OF 17 MEDLINE

- TI SU5416 is a potent and selective inhibitor of the vascular endothelial growth factor receptor (Flk-1/KDR) that inhibits tyrosine kinase catalysis, tumor vascularization, and growth of multiple tumor types.
- L6 ANSWER 10 OF 17 MEDLINE
- TI A follow-up study of patients hospitalized after minor head injury.
- L6 ANSWER 11 OF 17 MEDLINE
- TI A tumor-derived protein which provides T-cell costimulation through accessory cell activation.
- L6 ANSWER 12 OF 17 MEDLINE
- TI Ontogeny of tolerogen-responsive lymphocytes following neonatal inoculation of class II disparate semiallogeneic cells.
- L6 ANSWER 13 OF 17 MEDLINE
- TI The secreted tumor-associated antigen 90K is a potent immune stimulator.
- L6 ANSWER 14 OF 17 MEDLINE
- TI Self-help research and the public mental health system.
- L6 ANSWER 15 OF 17 MEDLINE
- TI In vitro suppression of cytotoxic T cell generation by lymphocytes from mice rendered neonatally tolerant of class II MHC alloantigens.
- L6 ANSWER 16 OF 17 MEDLINE
- TI Influence of I-E expression on induction of neonatal transplantation tolerance.
- L6 ANSWER 17 OF 17 MEDLINE
- TI I-E molecules and I-E-reactive T cells play a central role in neonatal H-2

tolerance.

=> DIS L6 1-10 IBIB ABS

THE ESTIMATED COST FOR THIS REQUEST IS 2.10 U.S. DOLLARS DO YOU WANT TO CONTINUE WITH THIS REQUEST? (Y)/N:Y

L6 ANSWER 1 OF 17 MEDLINE

ACCESSION NUMBER: 2000478714 MEDLINE

DOCUMENT NUMBER: 20483371 PubMed ID: 11030454

TITLE: Chronic neurobehavioural effects of mercury poisoning on a

group of Zulu chemical workers.

AUTHOR: Powell T J

CORPORATE SOURCE: West Berkshire Priority Care Services, NHS Trust, Reading,

UK.. trevorp@wbpcs-tr.anglox.nhs.uk

SOURCE: BRAIN INJURY, (2000 Sep) 14 (9) 797-814.

Journal code: 8710358. ISSN: 0269-9052.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200102

ENTRY DATE: Entered STN: 20010322

Last Updated on STN: 20010322

Entered Medline: 20010215

AB PRIMARY OBJECTIVE: To assess the nature and severity of reported neurobehavioural symptoms of mercury poisoning, in a group of Zulu chemical workers (n = 16), employed by a mercury processing plant, exposed

to neurotoxic levels of mercury, 5 years after exposure. RESEARCH DESIGN: A group-control design was adopted, where the exposed group was matched for age, sex, race, occupational and educational background. METHOD/PROCEDURES: Both groups were administered a specially selected battery of psychometric tests to measure neuropsychological functioning. OUTCOME AND RESULTS: The exposed group had significantly impaired short term verbal and spatial memory, impaired sustained and divided attention, and impaired motor speed. They also suffered from elevated clinical

of psychiatric symptomatology, including anxiety, depression and phobic avoidance, and neurological symptoms of tremor, weakness in the limbs,

excessive sweating. CONCLUSIONS: The exposed group suffered from varying degrees of permanent neuropsychological disability, which adversely affects their ability to work and be financially independent.

measures for monitoring cognitive symptoms are discussed.

ANSWER 2 OF 17 MEDLINE

ACCESSION NUMBER: 2000392640 MEDLINE

DOCUMENT NUMBER: 20287784 PubMed ID: 10828116

TITLE: Psychiatrists' referrals to self-help groups for people

with mood disorders.

AUTHOR: Powell T J; Silk K R; Albeck J H

CORPORATE SOURCE: School of Social Work, University of Michigan, Ann Arbor,

MI 48109-1106, USA.. tpowell@umich.edu

CONTRACT NUMBER: MH46399 (NIMH)

SOURCE: PSYCHIATRIC SERVICES, (2000 Jun) 51 (6) 809-11.

Journal code: 9502838. ISSN: 1075-2730.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200008

ENTRY DATE: Entered STN: 20000824

> Last Updated on STN: 20000824 Entered Medline: 20000817

AB The study examined psychiatrists' referrals to and support for participation in self-help groups by people with mood disorders. Massachusetts and Michigan psychiatrists with a special interest in patients with mood disorders were surveyed; the 278 respondents represented a 78 percent response rate. About three-fourths of the psychiatrists reported that they made referrals to and felt knowledgeable about self-help groups. However, less than half had self-help literature available or discussed self-help groups with their patients. Beliefs that a patient would gain a better understanding of the illness and would receive support after an episode of illness were positively related to support for self-help. Beliefs that the program was inappropriate and

that

it lacked professional oversight were negatively related.

ANSWER 3 OF 17 MEDLINE

ACCESSION NUMBER: 2000130287 MEDLINE

DOCUMENT NUMBER: 20130287 PubMed ID: 10662789

TITLE: A discrete subpopulation of dendritic cells transports

apoptotic intestinal epithelial cells to T cell areas of

mesenteric lymph nodes.

COMMENT: Comment in: J Exp Med. 2000 Feb 7;191(3):411-6 AUTHOR: Huang F P; Platt N; Wykes M; Major J R; Powell T J

; Jenkins C D; MacPherson G G

CORPORATE SOURCE: Sir William Dunn School of Pathology, University of

Oxford,

Oxford OX1 3RE, United Kingdom.

SOURCE: JOURNAL OF EXPERIMENTAL MEDICINE, (2000 Feb 7)

191 (3) 435-44.

Journal code: 2985109R. ISSN: 0022-1007.

PUB. COUNTRY:

United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200004

ENTRY DATE:

Entered STN: 20000427

Last Updated on STN: 20021231 Entered Medline: 20000418

AΒ This study identifies a dendritic cell (DC) subset that constitutively transports apoptotic intestinal epithelial cell remnants to T cell areas of mesenteric lymph nodes in vivo. Rat intestinal lymph contains two DC populations. Both populations have typical DC morphology, are major histocompatibility complex class II(hi), and express OX62, CD11c, and B7. CD4(+)/OX41(+) DCs are strong antigen-presenting cells (APCs).

CD4(-)/OX41(-) DCs are weak APCs and contain cytoplasmic apoptotic DNA, epithelial cell-restricted cytokeratins, and nonspecific esterase

(NSE)(+)

inclusions, not seen in OX41(+) DCs. Identical patterns of NSE electrophoretic variants exist in CD4(-)/OX41(-) DCs, intestinal epithelial cells, and mesenteric node DCs but not in other DC populations,

macrophages, or tissues. Terminal deoxynucleotidyl transferase-mediated dUTP-biotin nick-end labeling (TUNEL)-positive DCs and strongly NSE(+) DCs

are present in intestinal lamina propria. Peyer's patches and mesenteric but not other lymph nodes contain many strongly NSE(+) DCs in interfollicular and T cell areas. Similar DCs are seen in the ileum and

in

T cell areas of mesenteric nodes in gnotobiotic rats. These results show that a distinct DC subset constitutively endocytoses and transports apoptotic cells to T cell areas and suggest a role for these DCs in inducing and maintaining peripheral self-tolerance.

ANSWER 4 OF 17 MEDLINE

ACCESSION NUMBER: 2000108988 MEDLINE

DOCUMENT NUMBER: 20108988 PubMed ID: 10641012

TITLE: Derivation of temporary emergency exposure limits

(TEELs).

AUTHOR: Craig D K; Davis J S; Hansen D J; Petrocchi A J;

Powell T J; Tuccinardi T E Jr

CORPORATE SOURCE: Westinghouse Safety Management Solutions, Inc., Aiken, SC

29803, USA.

JOURNAL OF APPLIED TOXICOLOGY, (2000 Jan-Feb) 20 SOURCE:

(1) 11-20.

Journal code: 8109495. ISSN: 0260-437X.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 200003

ENTRY DATE: Entered STN: 20000407

> Last Updated on STN: 20000407 Entered Medline: 20000324

AB Short-term chemical concentration limits are used in a variety of applications, including emergency planning and response, hazard assessment

and safety analysis. Development of emergency response planning guidelines  $% \left( 1\right) =\left( 1\right) +\left( 1\right)$ 

(ERPGs) and acute exposure guidance levels (AEGLs) are predicated on this need. Unfortunately, the development of peer-reviewed community exposure limits for emergency planning cannot be done rapidly (relatively few ERPGs

or AEGLs are published each year). To be protective of Department of Energy (DOE) workers, on-site personnel and the adjacent general public, the DOE Subcommittee on Consequence Assessment and Protective Actions (SCAPA) has developed a methodology for deriving temporary emergency exposure limits (TEELs) to serve as temporary guidance until ERPGs or AEGLs can be developed. These TEELs are approximations to ERPGs to be

until peer-reviewed toxicology-based ERPGs, AEGL or equivalents can be developed. Originally, the TEEL method used only hierarchies of published concentration limits (e.g. PEL- or TLV-TWAs, -STELs or -Cs, and IDLHs) to provide estimated values approximating ERPGs. Published toxicity data (e.g. lc(50), lc(LO), ld(50) and ld(LO) for TEEL-3, and tc(LO) and td(LO) for TEEL-2) are included in the expanded method for deriving TEELs presented in this paper. The addition here of published toxicity data (in addition to the exposure limit hierarchy) enables TEELs to be developed for a much wider range of chemicals than before. Hierarchy-based values take precedence over toxicity-based values, and human toxicity data are used in preference to animal toxicity data. Subsequently, default assumptions based on statistical correlations of ERPGs at different

(e.g. ratios of ERPG-3s to ERPG-2s) are used to calculate TEELs where there are gaps in the data. Most required input data are available in the literature and on CD ROMs, so the required TEELs for a new chemical can be

developed quickly. The new TEEL hierarchy/toxicity methodology has been used to develop community exposure limits for over 1200 chemicals to date.

The new TEEL methodology enables emergency planners to develop useful approximations to peer-reviewed community exposure limits (such as the ERPGs) with a high degree of confidence. For definitions and acronyms,

Appendix

see

used

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L6 ANSWER 5 OF 17 MEDLINE

ACCESSION NUMBER: 2000050243 MEDLINE

DOCUMENT NUMBER: 20050243 PubMed ID: 10583160

TITLE: Growth inhibition of psoriatic keratinocytes by

quinazoline

tyrosine kinase inhibitors.

AUTHOR: Powell T J; Ben-Bassat H; Klein B Y; Chen H;

Shenoy N; McCollough J; Narog B; Gazit A; Harzstark Z; Chaouat M; Levitzki R; Tang C; McMahon J; Shawver L;

Levitzki A

CORPORATE SOURCE: SUGEN, Inc; Redwood City, CA 94063, USA.

SOURCE: BRITISH JOURNAL OF DERMATOLOGY, (1999 Nov) 141

(5) 802-10.

Journal code: 0004041. ISSN: 0007-0963.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

200005

ENTRY DATE:

Entered STN: 20000512

Last Updated on STN: 20000512 Entered Medline: 20000502

AB Psoriasis is characterized by hyperproliferation of keratinocytes associated with an inflammatory infiltrate in the epidermis. Among factors

which may be related to hyperplasia of psoriatic keratinocytes is the persistent autocrine stimulation of the epidermal growth factor receptor (EGFR) by transforming growth factor-alpha. Owing to the pivotal role of the EGFR in driving the growth of human psoriatic keratinocytes, we examined two selective inhibitors of EGFR kinase activity:

4-(3-bromophenylamino)-6, 7-dimethoxyquinazoline (AG1517/SU5271) and 4-(3-chlorophenylamino)-6, 7-dimethoxyquinazoline (AG1478) on psoriatic keratinocytes. SU5271 potently inhibits ligand-induced

autophosphorylation

of EGFR, and downstream signal transduction events, including DNA replication and cell cycle progression. SU5271, at micromolar concentrations, inhibited the proliferation of keratinocytes isolated

psoriatic lesions in excellent correlation with its EGFR kinase inhibitory

activity in these cells. Biologically active concentrations of SU5271 penetrated human cadaver skin, suggesting that this compound is a strong candidate as an antipsoriatic agent.

L6 ANSWER 6 OF 17 MEDLINE

ACCESSION NUMBER:

1999440218 MEDLINE

DOCUMENT NUMBER:

99440218 PubMed ID: 10510523

TITLE:

Recommended default methodology for analysis of airborne

exposures to mixtures of chemicals in emergencies.

AUTHOR:

Craig D K; Baskett R L; Davis J S; Dukes L; Hansen D J;

Petrocchi A'J; Powell T J; Sutherland P J;

Tuccinardi T E Jr

CORPORATE SOURCE:

Westinghouse Safety Management Solutions LLC, Aiken, South

Carolina, USA.

SOURCE:

APPLIED OCCUPATIONAL AND ENVIRONMENTAL HYGIENE, (1999

Sep) 14 (9) 609-17. Ref: 11

Journal code: 9103256. ISSN: 1047-322X.

PUB. COUNTRY:

ENGLAND: United Kingdom

DOCUMENT TYPE:

Journal; Article; (JOURNAL ARTICLE)

General Review; (REVIEW)

(REVIEW, TUTORIAL)

LANGUAGE:

English

FILE SEGMENT:

Priority Journals

ENTRY MONTH:

199910

ENTRY DATE:

Entered STN: 20000111

Last Updated on STN: 20000111

Entered Medline: 19991022

AB Emergency planning and hazard assessment of Department of Energy (DOE) facilities require consideration of potential exposures to mixtures of chemicals released to the atmosphere. Exposure to chemical mixtures may lead to additive, synergistic, or antagonistic health effects. In the past, the consequences of exposures to each chemical have been analyzed separately. This approach may not adequately protect the health of

exposed to mixtures. This article presents default recommendations for use

in emergency management and safety analysis within the DOE complex where

potential exists for releases of mixtures of chemicals. These recommendations were developed by the DOE Subcommittee on Consequence Assessment and Protective Actions (SCAPA). It is recommended that hazard indices (e.g., HIi = Ci/Limiti, where Ci is the concentration of chemical "i") be calculated for each chemical, and unless sufficient toxicological knowledge is available to indicate otherwise, that they be summed, that is, sigma i(n) = 1HIi = HI1 + HI2 + ... + HIn. A sum of 1.0 or less means the limits have not been exceeded. To facilitate application of these recommendations for analysis of exposures to specific mixtures, chemicals are classified according to their toxic consequences. This is done using health code numbers describing toxic effects by target organ for each chemical. This methodology has been applied to several potential releases of chemicals to compare the resulting hazard indices of a chemical

mixture

with those obtained when each chemical is treated independently. The methodology used and results obtained from analysis of one mixture are presented in this article. This article also demonstrates how health code numbers can be used to sum hazard indicés only for those chemicals that have the same toxic consequence.

L6 ANSWER 7 OF 17 MEDLINE

ACCESSION NUMBER: 1999372831 MEDLINE

DOCUMENT NUMBER: 99372831 PubMed ID: 10445659

TITLE: Attitudes of AA contact persons toward group participation

by persons with a mental illness.

AUTHOR: Meissen G; Powell T J; Wituk S A; Girrens K;

Arteaga S

CORPORATE SOURCE: Department of Psychology, Wichita State University, Kansas

67260, USA.. meissen@twsu.edu

SOURCE: PSYCHIATRIC SERVICES, (1999 Aug) 50 (8) 1079-81.

Journal code: 9502838. ISSN: 1075-2730.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199909

ENTRY DATE: Entered STN: 19990925

Last Updated on STN: 19990925 Entered Medline: 19990910

Alcoholics Anonymous groups are underused by persons with the dual diagnoses of mental illness and substance use disorder, and mental health professionals are cautious about referring them to AA because of fears that the AA group will discourage them from taking prescribed medication. The study assessed the attitudes of 125 AA contact persons about the participation of persons with mental illness. The majority had positive attitudes toward such persons, and 93 percent indicated that they should continue taking their medication. Fifty-four percent felt that participation in a group especially for persons with a dual diagnosis would be more desirable than in a traditional AA group. However, such groups are often not available.

L6 ANSWER 8 OF 17 MEDLINE

ACCESSION NUMBER: 1999111045 MEDLINE

DOCUMENT NUMBER: . 99111045 PubMed ID: 9815796

TITLE: Inhibition of platelet-derived growth factor-mediated

signal transduction and tumor growth by

N-[4-(trifluoromethyl)-phenyl]5-methylisoxazole-4-

carboxamide.

AUTHOR: Shawver L K; Schwartz D P; Mann E; Chen H; Tsai J; Chu L;

Taylorson L; Longhi M; Meredith S; Germain L; Jacobs J S;

Tang C; Ullrich A; Berens M E; Hersh E; McMahon G; Hirth K

P; Powell T J

CORPORATE SOURCE:

SUGEN, Inc., Redwood City, California 94063, USA.

SOURCE:

CLINICAL CANCER RESEARCH, (1997 Jul) 3 (7)

1167-77.

Journal code: 9502500. ISSN: 1078-0432.

PUB. COUNTRY:

United States

Journal; Article; (JOURNAL ARTICLE) DOCUMENT TYPE:

LANGUAGE:

English

FILE SEGMENT:

ENTRY MONTH:

Priority Journals

199902

ENTRY DATE:

Entered STN: 19990311

Last Updated on STN: 20000303 Entered Medline: 19990225

Many reports have cited coexpression of platelet-derived growth factor (PDGF) and its receptors by tumor cells or cells supporting tumor growth, suggesting both autocrine and paracrine mechanisms for PDGF-mediated

growth. We found that a small organic molecule, N-[4-(trifluoromethyl)phenyl] 5-methylisoxazole-4-carboxamide (SU101, leflunomide), inhibited PDGF-mediated signaling events, including

tyrosine phosphorylation, DNA synthesis, cell cycle progression, and cell proliferation. SU101 inhibited PDGF-stimulated tyrosine phosphorylation

PDGF receptor (PDGFR) beta in C6 (rat glioma) and NIH3T3 cells engineered to overexpress human PDGFRbeta (3T3-PDGFRbeta). SU101 blocked both PDGFand epidermal growth factor (EGF)-stimulated DNA synthesis. Previously, this compound was shown to inhibit pyrimidine biosynthesis by interfering with the enzymatic activity of dihydroorotate dehydrogenase. In the current study, EGF-stimulated DNA synthesis was restored by the addition of saturating quantities of uridine, whereas PDGF-induced DNA synthesis was not, suggesting that the compound demonstrated some selectivity for the PDGFR pathway that was independent of pyrimidine biosynthesis. Selectivity was further demonstrated by the ability of the compound to block the entry of PDGF-stimulated cells into the S phase of the cell cycle, without affecting cell cycle progression of EGF-stimulated cells. In cell growth assays, SU101 selectively inhibited the growth of PDGFRbeta-expressing cell lines more efficiently than it inhibited the growth of PDGFRbeta-negative cell lines. SU101 inhibited the s.c., i.p., and intracerebral growth of a panel of cell lines including cells from glioma, ovarian, and prostate origin. In contrast, SU101 failed to

the in vitro or s.c. growth of A431 and KB tumor cells, both of which express EGF receptor but not PDGFRbeta. SU101 also inhibited the growth

D1B and L1210 (murine leukemia) cells in syngeneic immunocompetent mice, without causing adverse effects on the immune response of the animals. In an i.p. model of tumor growth in syngeneic immunocompetent mice, SU101 prevented tumor growth and induced long-term survivors in animals implanted with 7TD1 (murine B-cell hybridoma) tumor cells. Because PDGFRbeta was detected on most of the tumor cell lines in which in vivo growth was inhibited by SU101, these data suggest that SU101 is an effective inhibitor of PDGF-driven tumor growth in vivo.

ANSWER 9 OF 17 MEDLINE

ACCESSION NUMBER: 1999107211 MEDLINE

DOCUMENT NUMBER: 99107211 PubMed ID: 9892193

SU5416 is a potent and selective inhibitor of the vascular TITLE:

endothelial growth factor receptor (Flk-1/KDR) that

inhibits tyrosine kinase catalysis, tumor vascularization,

and growth of multiple tumor types.

AUTHOR: Fong T A; Shawver L K; Sun L; Tang C; App H; Powell T

J; Kim Y H; Schreck R; Wang X; Risau W; Ullrich A;

Hirth K P; McMahon G

CORPORATE SOURCE: SUGEN, Inc., South San Francisco, California 94080, USA..

fong@progenitor.com

SOURCE: CANCER RESEARCH, (1999 Jan 1) 59 (1) 99-106.

Journal code: 2984705R. ISSN: 0008-5472.

PUB. COUNTRY: United States

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199902

ENTRY DATE: Entered STN: 19990216

Last Updated on STN: 20000303 Entered Medline: 19990204

AB SU5416, a novel synthetic compound, is a potent and selective inhibitor of

the Flk-1/KDR receptor tyrosine kinase that is presently under evaluation in Phase I clinical studies for the treatment of human cancers. SU5416

in Phase I clinical studies for the treatment of human cancers. SU541 was

shown to inhibit vascular endothelial growth factor-dependent mitogenesis of human endothelial cells without inhibiting the growth of a variety of tumor cells in vitro. In contrast, systemic administration of SU5416 at nontoxic doses in mice resulted in inhibition of subcutaneous tumor growth

of cells derived from various tissue origins. The antitumor effect of \$U5416 was accompanied by the appearance of pale white tumors that were resected from drug-treated animals, supporting the antiangiogenic property

of this agent. These findings support that pharmacological inhibition of the enzymatic activity of the vascular endothelial growth factor receptor represents a novel strategy for limiting the growth of a wide variety of tumor types.

L6 ANSWER 10 OF 17 MEDLINE

ACCESSION NUMBER: 96317074 MEDLINE

DOCUMENT NUMBER: 96317074 PubMed ID: 8743300

TITLE: A follow-up study of patients hospitalized after minor

head

injury.

AUTHOR: Powell T J; Collin C; Sutton K

CORPORATE SOURCE: Psychology Department, Erleigh Road Clinic, Reading, UK.

SOURCE: DISABILITY AND REHABILITATION, (1996 May) 18 (5)

231-7

Journal code: 9207179. ISSN: 0963-8288.

PUB. COUNTRY: ENGLAND: United Kingdom

DOCUMENT TYPE: Journal; Article; (JOURNAL ARTICLE)

LANGUAGE: English

FILE SEGMENT: Priority Journals

ENTRY MONTH: 199610

ENTRY DATE: Entered STN: 19961025

Last Updated on STN: 20000303 Entered Medline: 19961016

AB Minor head injury accounts for 95% of all head injury. In this study 62 patients, hospitalized after minor head injury, were assessed within 48 h.

and invited to attend for review and retesting 3 months later. Thirty-five  $\ensuremath{\text{3}}$ 

patients were followed up in this way and 11 more were interviewed over the telephone. There was significant improvement on all psychometric tests

between initial evaluation and follow-up. Between 51% and 86% reported troublesome late post-concussional symptoms, of which headaches and tiredness were the most frequently reported symptoms. Length of post-traumatic amnesia (PTA) was related to severity of symptoms. Clinical

levels of anxiety and stress were noted in approximately one-third of the whole group; 95% of the group had returned to work by 3 months with a mean

absence rate of 9.4 days. The therapeutic implications of these results are discussed.

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